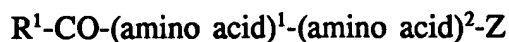


What is claimed is:

Sub.B. 1. ~~A reagent for preparing a scintigraphic imaging agent for imaging a site within a mammalian body, comprising a specific binding compound that is less than 10,000 daltons in molecular weight covalently linked to a radiolabel complexing moiety having formula:~~

I.



wherein (amino acid)¹ and (amino acid)² are each independently any primary α - or β -amino acid that does not comprise a thiol group;

Z is a thiol-containing moiety that is cysteine, homocysteine, isocysteine, penicillamine, 2-mercaptoethylamine or 3-mercaptopropylamine;

R¹ is lower (C¹-C⁴) alkyl or a covalent linkage to the specific binding compound;

wherein when Z is cysteine, homocysteine, isocysteine or penicillamine, the carbonyl group of said moiety is covalently linked to a hydroxyl group, a NR³R⁴ group, an amino acid or a peptide comprising 2 to 10 amino acids, and wherein R³ and R⁴ are each independently H or lower (C¹-C⁴) alkyl; or

II.



wherein Y is a thiol-containing moiety that is cysteine, homocysteine, isocysteine, penicillamine, 2-mercaptoacetate or 3-mercaptopropionate;

(amino acid)¹ and (amino acid)² are each independently any primary α - or β -amino acid that does not comprise a thiol group;

R² is H or lower (C¹-C⁴) alkyl or a covalent linkage to the specific binding compound;

wherein when Y is cysteine, homocysteine, isocysteine or penicillamine, the amino group of said moiety is covalently linked to -H, an amino acid or a peptide comprising 2 to 10 amino acids; and

wherein the radiolabel complexing moiety is covalently linked to the specific binding compound through R¹, R², a sidechain group of the sidechain of (amino acid)¹ or (amino acid)², or the amino or carboxyl group of cysteine, homocysteine, isocysteine or

penicillamine.

2. The reagent of Claim 1 wherein the radiolabel complexing moiety is selected from the group consisting of moieties having the formula:

-(amino acid)¹-(amino acid)²-(amino thiol),

and (mercaptocarboxylic acid)-(amino acid)¹-(amino acid)²-,

wherein (amino acid)¹ and (amino acid)² are each independently any primary α - or β -amino acid;

(amino thiol) is selected from the group consisting of cysteine, isocysteine, homocysteine, penicillamine, 2-mercaptoethylamine, and 3-mercaptopropylamine; and

(mercaptocarboxylic acid) is selected from the group consisting of cysteine, isocysteine, homocysteine, penicillamine, 2-mercaptoacetic acid, and 3-mercaptopropionic acid.

3. The reagent of Claim 2 wherein the radiolabel complexing moiety is selected

from the group consisting of moieties having the formula--Gly-Gly-Cys--or--Cys-Gly-Gly--

4. A composition of matter comprising the reagent according to Claim 1 selected from the group consisting of:

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCR.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRD.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRR.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKKK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKDK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.D.Orn.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.D.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KKC.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KRC.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.RRC.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KKCK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GRCK.amide)

cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GKCR.amide)

CH₂CO.Y_D.Apc.GDCGGC_{Ac}GC_{Ac}GGC.amide
 CH₂CO.Y_D.Apc.GDCGGC_{Ac}GC_{Ac}GGCG.amide
 CH₂CO.Y_D.Apc.GDCGGSSGGCG.amide
 CH₂CO.Y_D.Apc.GDCGGCG.amide
 5 GRGDGGC
 GLFCGC.amide
 GRGDGGGGC
 F_DFYW_DKTFTGGC.amide
 acetyl.CGGY.(CH₂)₄-piperidine
 10 β-glucan-(=NNHCO.(CH₂)₆CO.)GGC.amide

Sub. B2
~~5. A reagent according to Claim 1 wherein the specific binding compound is a specific binding peptide comprising 4 to 100 amino acids.~~

~~6. The reagent of Claim 1 wherein the specific binding peptide and radiolabel binding moiety are covalently linked through one or more amino acids.~~

7. A scintigraphic imaging agent comprising the reagent according to Claim 1 wherein the radiolabel binding moiety is bound to a radiolabel.

8. The reagent of Claim 7 wherein the radiolabel is technetium-99m.

9. ~~The reagent of Claim 1 wherein the reagent further comprises a polyvalent linking moiety covalently linked to a multiplicity of specific binding compounds and also covalently linked to a multiplicity of radiolabel-complexing moieties to comprise a reagent for preparing a multimeric polyvalent scintigraphic imaging agent, wherein the molecular weight of the multimeric polyvalent scintigraphic imaging agent is less than about 20,000 daltons.~~

~~10. The reagent of Claim 9 wherein the polyvalent linking moiety is bis-succinimethylmethylether, 4-(2,2-dimethylacetyl)benzoic acid, tris(succinimidylethyl) amine, 4-(O-CH₂CO-Gly-Gly-Cys.amide)acetophenone, bis-succinimidohexane, tris(2-chloroacetamidoethyl)amine, and 1,2-bis-[2-(chloroacetamido)ethoxy]ethane or a derivative thereof.~~

11. A complex formed by reacting the reagent of Claim 1 with technetium-99m in the presence of a reducing agent.

12. The complex of Claim 11, wherein the reducing agent is selected from the group consisting of a dithionite ion, a stannous ion and a ferrous ion.

13. A complex formed by labeling the reagent of Claim 1 with technetium-99m by ligand exchange of a prereduced technetium-99m complex.

14. A kit for preparing a radiopharmaceutical preparation, said kit comprising a sealed vial containing a predetermined quantity of the reagent of Claim 1 and a sufficient amount of reducing agent to label the reagent with technetium-99m.

15. A method for labeling a reagent according to Claim 1 comprising reacting the reagent with technetium-99m in the presence of a reducing agent.

16. The method of Claim 15, wherein the reducing agent is selected from the group consisting of a dithionite ion, a stannous ion and a ferrous ion.

17. A method for imaging a site within a mammalian body comprising administering an effective diagnostic amount of the reagent of Claim 2 and detecting a radioactive signal from the technetium-99m localized at the site.

18. A composition of matter having formula:

Sub B
15 cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCK.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCR.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRD.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRK.amide)
20 cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCRR.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKK.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKKK.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGCKDK.amide)
25 cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.D.Orn.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GGC.Orn.D.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KKC.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KRC.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.RRC.amide)
30 cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.KKCK.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GRCK.amide)
cyclo(N-methyl)FYW_DKV.Hcy.(CH₂CO.GKCR.amide)
CH₂CO.Y_D.Apc.GDCGGC_{Ac}GC_{Ac}GGC.amide
CH₂CO.Y_D.Apc.GDCGGC_{Ac}GC_{Ac}GGCG.amide
35 CH₂CO.Y_D.Apc.GDCGGSSGGCG.amide
CH₂CO.Y_D.Apc.GDCGGCG.amide
GRGDGGC
GLFCGC.amide

GRGDGGGGC

F_DFYW_DKTFTGGC.amide

aceryl.CGGY.(CH₂)₄-piperidine

or

β-glucan-(=NNHCO.(CH₂)₃CO.)GGC.amide

19. The reagent of Claim 1 wherein the specific binding peptide is comprised of linear or cyclic peptides.

20. The reagent of Claim 1 wherein the imaged site within a mammalian body is a thrombus site.

21. The reagent of Claim 1 wherein the imaged site within a mammalian body is a site of an infection.

22. A composition of matter according to Claim 18 that is radiolabeled with technetium-99m.

23. An article of manufacture comprising a sealed vial containing a predetermined quantity of the composition of matter of Claim 18 and a sufficient amount of reducing agent to label the composition with technetium-99m.